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(BAYESIAN INFERENCE POSTERIOR DISTRIBUTIONS PREDICTIVE ANALYSIS PRIOR DISTRIBUTIONS)

LARA J. WOLFSON

EMPIRICAL BIVARIATE QUANTILE-PARTITIONED DISTRIBUTION

Epidemiologists sometimes collect bivariate continuous data on a sample of individuals, calculate the empirical quantiles* of the marginal data, and then partition the original data into two-way contingency tables* with row and column categories defined by these values [16]. For example, Pietinen et al. [13, 14] conducted an extensive study on Finnish men aged 55-69 to test the reproducibility and validity of dietary measurement techniques. As part of this study, the vitamin E intake of 157 men was measured by a prospective food record diary (X) and a retrospective food use questionnaire (Y). In turn, these bivariate measurements were partitioned into categories defined by the empirical quintiles of the X and Y variables to create Table 1 [4]. The term empirical bivariate quantile-partitioned (EBQP) distribution describes the distribution of counts in such tables [3].

Because the original data are partitioned by the empirical quintiles, the marginal totals of

Table 1 EBQP Table for the Vitamin E Data Partitioned by Empirical Quintiles^a

Food Record Quintiles	Food Use Questionnaire Quintiles					
	l (low)	2	3	4	5 (high)	Total
1 (low)	13	13	4	1	0	31
2	9	9	7	4	2	31
3	7	5	7	7	6	32
4	2	2	12	9	6	31
5 (high)	0	2	2	10	18	32
Total	31	31	32	31	32	157

^aCreated from original data in a study by Pietinen et al. [13], adapted from ref. [4].

the rows and column are fixed at 31 or 32 observations, even though the interior counts of the table are still random. For example, 12 individuals fell in both the fourth quintile of the food record measurements and the third quintile of the food use questionnaire measurements.

The special case of 2×2 EBOP tables partitioned by empirical medians has received considerable study. In 1899, Sheppard [15] proposed studying the agreement between two bivariate continuous measurements by constructing such tables. Blomqvist [1] discussed a measure of agreement*, q, adapted from Mosteller [12] for such tables. This measure, called the medial correlation [10] or the quadrant measure [11], is algebraically equivalent to Kendall's tau* in 2×2 tables*. In order to study the distribution of such statistics, Blomqvist developed an exact theory and derived an asymptotic variance for 2×2 tables under certain regularity conditions, amended by Konijn [10]. Elandt [6, 7] discussed the power of Blomqvist's exact test of independence under bivariate normal alternatives.

Borkowf et al. [3] derived the asymptotic normal distribution theory for the cell proportions in $r \times c$ EBQP tables, and gave a special formula for the 2×2 case. They found, to their surprise, that Blomqvist's asymptotic result for 2×2 EBQP tables was correct only in special cases. Measures of agreement calculated from $r \times c$ EBQP tables, such as kappa* or weighted kappa, are functions of the cell proportions in these tables. Hence, the asymptotic normal distribution of these measures can be derived from the asymptotic joint normal distribution of the cell proportions, and the variances of these measures can be calculated by the delta method*.

The EBQP method of constructing tables and the corresponding method of inference for measures calculated from these tables are both nonparametric*. By contrast, if one is willing to assume that the original bivariate measurements come from a particular parametric family, one can estimate the parameters that determine the shape of the underlying distribution, calculate the parametric estimates of the population quantiles, and then use these values to estimate the

expected counts in tables analogous to Table 1. Borkowf and Gail [5] developed the parametric method in order to study the asymptotic relative efficiency* of EBQP methods and to improve the precision of estimates of measures of agreement.

Here we review the asymptotic distribution theory for $r \times c$ EBQP tables, discuss the efficiency of EBQP methods compared to parametric methods, and illustrate the use of these methods with an example.

NOTATION AND ASSUMPTIONS

Let the bivariate continuous sample $\{(X_k, Y_k)\}$ (k = 1, 2, ..., t) be i.i.d. from the distribution F. Let F(x, y) have marginal distributions G(x) and H(y) and conditional distributions G(x|y) and H(y|x). Also, let $\hat{F}(x, y)$, $\hat{G}(x)$, and $\hat{H}(y)$ denote the corresponding right-continuous empirical distribution functions (see EDF STATISTICS).

Next, let $\{\gamma_i\}$ ($i=0,1,\ldots,r$) and $\{\eta_j\}$ ($i=0,1,\ldots,c$) denote two increasing sets of cumulative marginal proportions such that $\gamma_0=\eta_0=0$ and $\gamma_r=\eta_c=1$. For example, for quintiles, r=c=5, $\gamma_i=i/5$, and $\eta_j=j/5$. In turn, define the population quantiles $\xi_i=G^{-1}(\gamma_i)$ and $\psi_j=H^{-1}(\eta_j)$, the empirical quantiles $u_i=\inf\{u:\gamma_i\leqslant \hat{G}(u)\}$ and $v_j=\inf\{v:\eta_j\leqslant \hat{H}(v)\}$, the conditional proportions $\gamma_{i|j}=G(\xi_i|\psi_j)$ and $\eta_{j|i}=H(\psi_j|\xi_i)$, and the cumulative proportions $\phi_{ij}=F(\xi_i,\psi_i)$.

It is assumed that g(x) = G'(x) and h(y) = H'(y) exist and are positive at the selected population quantiles, so $\xi_i = G^{-1}(\gamma_i)$ and $\psi_j = H^{-1}(\eta_j)$ are uniquely defined. It is also assumed that F(x, y) is differentiable as a function of (x, y) at each (ξ_i, ψ_j) .

Then, the proportion of counts in the (i, j)th category defined by $u_{i-1} < x \le u_i$ and $v_{j-1} < y \le v_j$ is

$$p_{ij} = \hat{F}(u_i, v_j) - \hat{F}(u_{i-1}, v_j) - \hat{F}(u_i, v_{j-1}) + \hat{F}(u_{i-1}, v_{j-1}).$$
(1)

Thus, the cell counts in the $r \times c$ EBQP table are given by $\{p_{ij}t\}$. As $t \to \infty$, each empirical proportion p_{ij} tends to the asymptotic

parametric methods, the corresponding point estimates and SEs are $\hat{\kappa} = 0.242 \pm 0.027$, $\hat{\kappa}_{w} = 0.627 \pm 0.042$, and $\hat{\alpha}_{111} = 0.551 \pm 0.031$.

Under EBQP methods, $\hat{\kappa}_w$ depends on all the cells of the table, whereas $\hat{\kappa}$ depends only on the diagonal cells and $\hat{\alpha}_{1|1}$ depends only on the (1,1) cell. By contrast, parametric methods use information from all of the underlying data through the parameter estimates (in this case, the sample correlation) to estimate these measures. Thus, it is not surprising that EBQP methods are most efficient for estimating κ_w and least efficient for estimating κ and $\alpha_{1|1}$, compared to parametric methods.

DISCUSSION

The EBQP theory presented here yields results that differ from those appropriate for statistics calculated from tables with the multinomial distribution*. Such tables are obtained if the original data are partitioned by the population quantiles rather than the empirical quantiles. Fleiss et al. [9] derived the asymptotic variances of kappa and weighted kappa calculated from multinomial tables, but these variances, though correct for multinomial tables, differ from the correct variances for statistics calculated from EBQP tables [3].

As the above example illustrates, EBQP methods can be inefficient compared to parametric methods for estimating certain measures of agreement. For BVN data, EBQP estimates of κ and $\alpha_{1|1}$ are less than 41% efficient for a range of correlations and table dimensions studied in ref. [5]. By contrast, EBQP estimates of κ_w become increasingly efficient as the table dimensions increase, for moderate correlations. The increased efficiency of parametric methods, however, comes at the risk of bias from misspecifying the parametric model [5].

EBQP methods are most helpful when used in conjunction with methods that directly examine the original bivariate continuous data. A full discussion of EBQP and parametric methods in the epidemiological context appears in ref. [4]. Sample computer code for performing EBQP and parametric calculations appears in ref. [2] and, at the time of this writing, can be obtained from the first author.

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proportion

$$\pi_{ij} = F(\xi_i, \psi_j) - F(\xi_{i-1}, \psi_j) - F(\xi_i, \psi_{j-1}) + F(\xi_{i-1}, \psi_{j-1}).$$
(2)

The empirical marginal proportions p_{i+} and p_{+j} are constant given t, while the asymptotic marginal proportions $\pi_{i+} = \gamma_i - \gamma_{i-1}$ and $\pi_{+j} = \eta_j - \eta_{j-1}$ are constant for all t.

ASYMPTOTIC THEORY

Utilizing (1), the asymptotic joint normal distribution, expectations, and covariances of the $\{p_{ij}\}$ can be derived from those of the $\{\hat{F}(u_i, v_i)\}$.

One can approximate $\hat{F}(u_i, v_j)$ in terms of $\hat{F}(\xi_i, \psi_j)$, $\hat{G}(\xi_i)$, and $\hat{H}(\psi_j)$. Define the vectors $\lambda'_{ij} = (1, -\eta_{j|i}, -\gamma_{i|j}), \mu'_{ij} = (0, \gamma_i, \eta_j)$, and $w'_{ij} = (\hat{F}(\xi_i, \psi_j), \hat{G}(\xi_i), \hat{H}(\psi_j))$. Then [3]

$$\hat{F}(u_i, v_j) = \lambda'_{ij}(w_{ij} - \mu_{ij}) + o_p(t^{-1/2}).$$
(3)

Since $t^{1/2}(\hat{F} - F, \hat{G} - G, \hat{H} - H)$ evaluated at the population quantiles has an asymptotic joint normal distribution, (3) implies that each $t^{1/2}\hat{F}(u_i, v_j)$ tends to normality, and moreover the vector $t^{1/2}\{\hat{F}(u_i, v_j)\}_{ij}$ jointly tends to normality.

Let $m = \min\{i, k\}$ and $n = \min\{j, l\}$. Then [3] for every sample size t,

$$E[\hat{F}(\xi_i, \psi_j)] = \phi_{ij}, \qquad (4)$$

$$Cov[t^{1/2}\hat{F}(\xi_i, \psi_j), t^{1/2}\hat{F}(\xi_k, \psi_1)] = \phi_{mn} - \phi_{ij}\phi_{kl};$$
(5)

also $G(x) = F(x, \infty)$ and $H(y) = F(\infty, y)$, so $\gamma_i = \phi_{ic}$ and $\eta_j = \phi_{rj}$. Thus, (5) implies that $Cov[t^{1/2}w_{ij}, t^{1/2}w_{kl}]$

$$=\begin{bmatrix} \phi_{mn} - \phi_{ij}\phi_{kl} & \phi_{mj} - \phi_{ij}\gamma_k & \phi_{in} - \phi_{ij}\eta_l \\ \phi_{ml} - \gamma_i\phi_{kl} & \gamma_m - \gamma_i\gamma_k & \phi_{il} - \gamma_i\eta_l \\ \phi_{kn} - \eta_j\phi_{kl} & \phi_{kj} - \eta_j\gamma_k & \eta_n - \eta_j\eta_l \end{bmatrix}$$

$$\equiv \Omega_{ijkl}. (6)$$

It follows from (4), (5), and (6) that $t^{1/2}[\hat{F}(u_i, v_j) - \phi_{ij}]$ and $t^{1/2}[\hat{F}(u_k, v_l) - \phi_{kl}]$ are jointly asymptotic normal with mean zero and covariance

$$\lambda'_{ij}\Omega_{ijkl} \lambda_{kl}$$
. (7)

In particular, the asymptotic variance of $t^{1/2}[\hat{F}(u_i,v_j)-\phi_{ij}]$ can be written without matrix notation as

$$\phi_{ij}(1 - \phi_{ij}) + \eta_{j|i}^{2} \gamma_{i}(1 - \gamma_{i}) + \gamma_{i|j}^{2} \eta_{j}(1 - \eta_{j})$$

$$- 2\eta_{j|i} \phi_{ij}(1 - \gamma_{i}) - 2\gamma_{i|j} \phi_{ij}(1 - \eta_{j})$$

$$+ 2\eta_{j|i} \gamma_{i|j}(\phi_{ij} - \gamma_{i}\eta_{j}). \quad (8)$$

In 2×2 tables partitioned by empirical medians, (8) reduces to Blomqvist's result only when $\gamma_{1|1} = \eta_{1|1} = \frac{1}{2}$. This condition holds not only under independence, but also for the bivariate normal (BVN) distribution with nonzero correlation, for instance.

For most applications, the variances and covariances involve so many terms that it is essential to use matrix notation and computer calculations. For a discussion of parameter estimation and confidence interval construction, see ref. [3].

ANALYSIS OF EXAMPLE

An analysis of the vitamin E data illustrates the use of EBQP methods in epidemiology [4]. To construct Table 1, several ties in the vitamin E data were broken by adding tiny random errors to the original data, but the manner in which these ties were broken had only a small effect on the resulting table. Consider the following three measures of agreement: kappa (κ), weighted kappa (κ_w) with quadratic weights [8], and row proportions [16] (the proportion of observations that fall in specified columns of a table, given that they fall in a specified row, e.g., $\alpha_{1|1} = \pi_{1|1}/\pi_{1+1}$). These measures and their variances can be estimated using the methods in ref. [3], which require not only the cell counts but also the original bivariate data. Using EBQP methods, the point estimates and standard errors (SEs) for these measures of agreement are $\hat{\kappa} = 0.196 \pm 0.056$, $\hat{\kappa}_w = 0.631 \pm 0.052$, and $\hat{\alpha}_{1|1} = 0.419 \pm 0.076$.

Normal probability plots and scatter plots of the log-transformed vitamin E data suggest that the original data are consistent with an underlying bivariate lognormal distribution. Hence, the log-transformed data are consistent with a BVN distribution with correlation $\hat{\rho} = 0.681$. Using

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(CONTINGENCY TABLES
EDF STATISTICS
KAPPA COEFFICIENT
MEASURES OF AGREEMENT
NICKED SQUARE DISTRIBUTION
QUANTILES
TWO-BY-TWO TABLES)

CRAIG B. BORKOWF MITCHELL H. GAIL

EPIDEMICS AMONG INTRAVENOUS DRUG USERS

The usual model for an epidemic*, as outlined in Bailey [1], assumes that the spread of infection depends on the law of mass action. If, for example, we have a deterministic model in which x(t) susceptibles (individuals liable to infection) mix with y(t) infectives (infected individuals who can transmit the infection) in a

population at time $t \ge 0$, then in the absence of a removal mechanism, the number of infectives will increase at a rate proportional to the product x(t)y(t).

While such a model provides a reasonable representation of the spread of infection by contacts between individuals, it is not entirely suitable for infections spread through the exchange of needles among intravenous drug users (IVDUs). Several authors, among them Doll [2], have noted the rapid rise of AIDS cases in the USA and other countries due to the exchange of infected needles among IVDUs. While most studies of epidemics among IVDUs have concentrated on HIV and AIDS, the models outlined below apply to any infection transmitted by blood such as, for example, hepatitis.

Different models have been developed for epidemics among IVDUs, of which two are typical. Firstly, Kaplan [3] has studied a deterministic model based on principles derived from the mass action model. He first finds an equation for the time-dependent probability that an IVDU injects with an infected needle at time $t \ge 0$, and then derives a differential equation involving a mass action factor for the fraction of infected IVDUs at time t in a shooting gallery. He proceeds to analyze various factors such as the sharing rates of injection equipment and their heterogeneity, the mean duration of injection equipment sharing, and the effect of cleaning injection equipment after use. Kaplan provides graphs to illustrate aspects of the model.

Secondly, Gani and Yakowitz [4] have considered a stochastic model where a group of n IVDUs consisting of X(t) susceptibles and Y(t) infectives, with X(t) + Y(t) = n, meets regularly at times t = 0, 1, 2, ..., to inject with drugs. The probability of creating new infectives at such a meeting was derived by Gani [5] in the context of an occupancy problem*. The susceptibles are thought of as cells, while the infective IVDUs who pass their infected needles to them are balls placed in these cells. Assuming the random allocation of infected needles among the susceptibles, and using the occupancy probabilities, one can characterize X(t)as a nonincreasing homogeneous Markov chain whose transition probabilities can be found. It is then easy to calculate the time until the

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